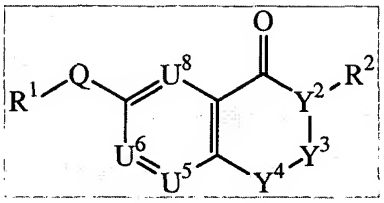
CLAIMS

What is claimed is:

1. A compound of Formula I



I

or a pharmaceutically acceptable salt thereof,

wherein:

R^1 is independently selected from:

C_5 or C_6 cycloalkyl-(C_1-C_8 alkylenyl);

Substituted C_5 or C_6 cycloalkyl-(C_1-C_8 alkylenyl);

C_8-C_{10} bicycloalkyl-(C_1-C_8 alkylenyl);

Substituted C_8-C_{10} bicycloalkyl-(C_1-C_8 alkylenyl);

5- or 6-membered heterocycloalkyl-(C_1-C_8 alkylenyl);

Substituted 5- or 6-membered heterocycloalkyl-(C_1-C_8 alkylenyl);

8- to 10-membered heterobicycloalkyl-(C_1-C_8 alkylenyl);

Substituted 8- to 10-membered heterobicycloalkyl-(C_1-C_8 alkylenyl);

Phenyl-(C_1-C_8 alkylenyl);

Substituted phenyl-(C_1-C_8 alkylenyl);

Naphthyl-(C_1-C_8 alkylenyl);

Substituted naphthyl-(C_1-C_8 alkylenyl);

5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl);

Substituted 5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl);

8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl);

Substituted 8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl);

Phenyl;

Substituted phenyl;

Naphthyl;

Substituted naphthyl;

5- or 6-membered heteroaryl;

Substituted 5- or 6-membered heteroaryl;

8- to 10-membered heterobiaryl; and

Substituted 8- to 10-membered heterobiaryl;

R^2 is independently selected from:

H;

C_1-C_6 alkyl;

Phenyl-(C_1-C_8 alkylenyl);

Substituted phenyl-(C_1-C_8 alkylenyl);

Naphthyl-(C_1-C_8 alkylenyl);

Substituted naphthyl-(C_1-C_8 alkylenyl);

5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl);

Substituted 5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl);

8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl);

Substituted 8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl);

Phenyl-O-(C_1-C_8 alkylenyl);

Substituted phenyl-O-(C_1-C_8 alkylenyl);

Phenyl-S-(C_1-C_8 alkylenyl);

Substituted phenyl-S-(C_1-C_8 alkylenyl);

Phenyl-S(O)-(C_1-C_8 alkylenyl);

Substituted phenyl-S(O)-(C_1-C_8 alkylenyl);

Phenyl-$S(O)_2$-(C_1-C_8 alkylenyl); and

Substituted phenyl-$S(O)_2$-(C_1-C_8 alkylenyl);

Each substituted R^1 and R^2 group contains from 1 to 4 substituents, each independently on a carbon or nitrogen atom, independently selected from:

C_1-C_6 alkyl;

CN;

CF_3;

HO;

(C_1-C_6 alkyl)-O;

$(C_1\text{-}C_6$ alkyl)-$S(O)_2$;

H_2N;

$(C_1\text{-}C_6$ alkyl)-N(H);

$(C_1\text{-}C_6$ alkyl)$_2$-N;

$(C_1\text{-}C_6$ alkyl)-C(O)O-$(C_1\text{-}C_8$ alkylenyl)$_m$;

$(C_1\text{-}C_6$ alkyl)-C(O)O-(1- to 8-membered heteroalkylenyl)$_m$;

$(C_1\text{-}C_6$ alkyl)-C(O)N(H)-$(C_1\text{-}C_8$ alkylenyl)$_m$;

$(C_1\text{-}C_6$ alkyl)-C(O)N(H)-(1- to 8-membered heteroalkylenyl)$_m$;

$H_2NS(O)_2$-$(C_1\text{-}C_8$ alkylenyl);

$(C_1\text{-}C_6$ alkyl)-N(H)$S(O)_2$-$(C_1\text{-}C_8$ alkylenyl)$_m$;

$(C_1\text{-}C_6$ alkyl)$_2$-$NS(O)_2$-$(C_1\text{-}C_8$ alkylenyl)$_m$;

3- to 6-membered heterocycloalkyl-$(G)_m$;

Substituted 3- to 6-membered heterocycloalkyl-$(G)_m$;

5- or 6-membered heteroaryl-$(G)_m$;

Substituted 5- or 6-membered heteroaryl-$(G)_m$;

$(C_1\text{-}C_6$ alkyl)-$S(O)_2$-N(H)-C(O)-$(C_1\text{-}C_8$ alkylenyl)$_m$; and

$(C_1\text{-}C_6$ alkyl)-C(O)-N(H)-$S(O)_2$-$(C_1\text{-}C_8$ alkylenyl)$_m$;

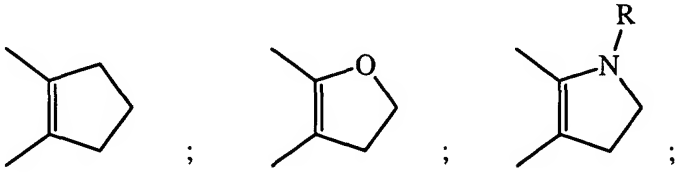
wherein each substituent on a carbon atom may further be independently selected from:

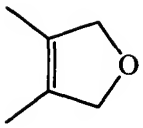
Halo; and

HO_2C;

wherein 2 substituents may be taken together with a carbon atom to which they are both bonded to form the group C(=O);

wherein two adjacent, substantially sp^2 carbon atoms may be taken together with a diradical substituent to form a cyclic diradical selected from:

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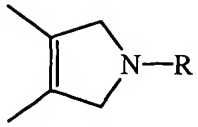
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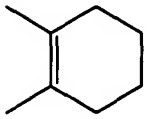
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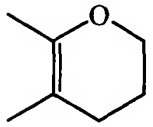
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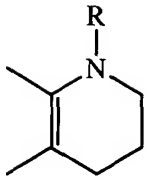
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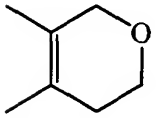
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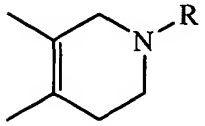
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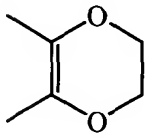
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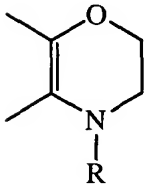
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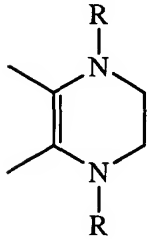
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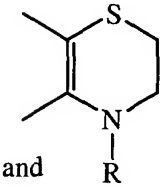
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R is H or C_1-C_6 alkyl;

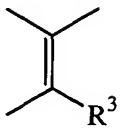
G is CH_2; O, S, S(O); or $S(O)_2$;

m is an integer of 0 or 1;

Y^2 is N;

Y^3 is CH_2; or

Y^2 and Y^3 are taken together to form the diradical group: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Y^4 is O or N-R^5, wherein R^5 is H or C_1-C_6 alkyl;

U^5, U^6, and U^8 are each C(H); or

1 of U^5, U^6, and U^8 is C-R^4 or N and the other 2 of U^5, U^6, and U^8 are each C(H);

R^3 and R^4 are independently selected from the groups:

H;

F;

Cl;

CH_3;

CH_3O;

$CH=CH_2$;

HO;

CF_3; and

CN;

Q is selected from:

OC(O);

$CH(R^6)C(O)$;

$OC(NR^6)$;

$CH(R^6)C(NR^6)$;

$N(R^6)C(O)$;

$N(R^6)C(S)$;

$N(R^6)C(NR^6)$;

$N(R^6)CH_2$;

SC(O);

$CH(R^6)C(S)$;

$SC(NR^6)$;

trans-(H)C=C(H);

cis-(H)C=C(H);

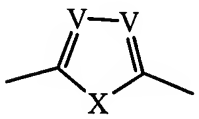
C≡C;

$CH_2C≡C$;

$C≡CCH_2$;

$CF_2C≡C$; and

$C≡CCF_2$;



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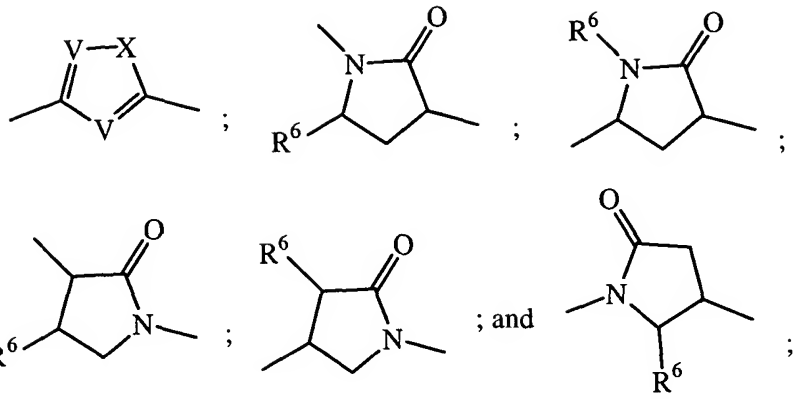


Each R^6 independently is H, C_1-C_6 alkyl, C_3-C_6 cycloalkyl; 3- to 6-membered heterocycloalkyl; phenyl; benzyl; or 5- or 6-membered heteroaryl;

X is O, S, N(H), or N(C_1-C_6 alkyl);

Each V is independently C(H) or N;

wherein each C_8-C_{10} bicycloalkyl is a bicyclic carbocyclic ring that contains 8-, 9-, or 10-member carbon atoms which are 5,5-fused, 6,5-fused, or 6,6-fused bicyclic rings, respectively, and wherein the ring is saturated or optionally contains one carbon-carbon double bond;

wherein each 8- to 10-membered heterobicycloalkyl is a bicyclic ring that contains carbon atoms and from 1 to 4 heteroatoms independently selected from 2 O, 1 S, 1 S(O), 1 $S(O)_2$, 1 N, 4 N(H), and 4 N(C_1-C_6 alkyl), and wherein when two O atoms or one O atom and one S atom are present, the two O atoms or one O atom and one S atom are not bonded to each other, and wherein the ring is saturated or optionally contains one carbon-carbon or carbon-nitrogen double bond, and wherein the heterobicycloalkyl is a 5,5-fused, 6,5-fused, or 6,6-fused bicyclic ring, respectively,

wherein each heterocycloalkyl is a ring that contains carbon atoms and from 1 to 4 heteroatoms independently selected from 2 O, 1 S, 1 S(O), 1 $S(O)_2$, 1 N, 4 N(H), and 4 N(C_1-C_6 alkyl), and wherein when two O atoms or one O atom and one S atom are present, the two O atoms or one O atom and one S atom are not bonded to each other, and wherein the ring is saturated or optionally contains one carbon-carbon or carbon-nitrogen double bond;

wherein each 5-membered heteroaryl contains carbon atoms and from 1 to 4 heteroatoms independently selected from 1 O, 1 S, 1 N(H), 1 N(C_1-C_6 alkyl), and 4 N, and each 6-membered heteroaryl contains carbon atoms and 1 or 2 heteroatoms independently selected from N, N(H), and N(C_1-C_6 alkyl), and 5- and 6-membered heteroaryl are monocyclic rings;

wherein each heterobiaryl contains carbon atoms and from 1 to 4 heteroatoms independently selected from 1 O, 1 S, 1 N(H), 1 N(C_1-C_6 alkyl), and 4 N, and where the 8-, 9-, and 10-membered heterobiaryl are 5,5-fused, 6,5-fused, and 6,6-fused bicyclic rings, respectively, and wherein at least 1 of the 2 fused rings of a bicyclic ring is aromatic, and wherein when the O and S atoms both are present, the O and S atoms are not bonded to each other;

wherein with any (C_1-C_6 alkyl)$_2$-N group, the C_1-C_6 alkyl groups may be optionally taken together with the nitrogen atom to which they are attached to form a 5- or 6-membered heterocycloalkyl; and

wherein each group and each substituent recited above is independently selected.

2. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein U^5, U^6, and U^8 are each C(H).

3. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein one of U^5, U^6, and U^8 is C-R^4 and the other two of U^5, U^6, and U^8 are each C(H).

4. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein one of U^5, U^6, and U^8 is N and the other two of U^5, U^6, and U^8 are each C(H).

5. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein Q is N(R^6)C(O).

6. The compound according to Claim 1, or a pharmaceutically acceptable salt thereof, wherein Q is C≡C

7. The compound according to any one of Claims 1 to 6, or a pharmaceutically acceptable salt thereof, wherein R^1 is independently selected from:

Phenyl-(C_1-C_8 alkylenyl);

Substituted phenyl-(C_1-C_8 alkylenyl);

5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl);

Substituted 5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl);

8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl); and

Substituted 8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl); and

R^2 is independently selected from:

Phenyl-(C_1-C_8 alkylenyl)$_m$;

Substituted phenyl-(C_1-C_8 alkylenyl)$_m$;

5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl)$_m$;

Substituted 5- or 6-membered heteroaryl-(C_1-C_8 alkylenyl)$_m$;

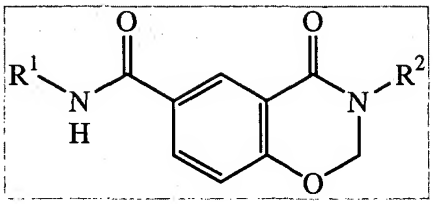
8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl)$_m$; and

Substituted 8- to 10-membered heterobiaryl-(C_1-C_8 alkylenyl)$_m$;

wherein m is an integer of 0 or 1; and

wherein each group and each substituent is independently selected.

8. The compound of Claim 1 of Formula II, IV, V, or VII



II

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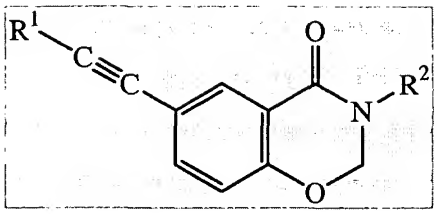

IV

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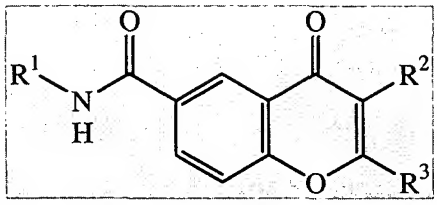

V

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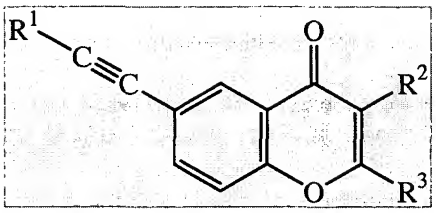

VII

9. The compound according to Claim 8 of Formula II selected from:

4-(6-Benzylcarbamoyl-4-oxo-4H-benzo[e][1,3]oxazin-3-ylmethyl)-benzoic acid;

4-[6-(4-Fluoro-benzyl)-carbamoyl-4-oxo-4H-benzo[e][1,3]oxazin-3-ylmethyl]-benzoic acid;

3-(4-Fluoro-benzyl)-4-oxo-3,4-dihydro-2H-benzo[e][1,3]oxazine-6-carboxylic acid benzylamide; and

3-(4-Fluoro-benzyl)-4-oxo-3,4-dihydro-2H-benzo[e][1,3]oxazine-6-carboxylic acid 4-methoxy-benzylamide; or

a pharmaceutically acceptable salt thereof.

10. The compound according to Claim 8 of Formula IV selected from:

4-[4-Oxo-6-(3-phenyl-prop-1-ynyl)-4H-benzo[e][1,3]oxazin-3-ylmethyl]-benzoic acid;

4-{6-[3-(4-Fluoro-phenyl)-prop-1-ynyl]-4-oxo-4H-benzo[e][1,3]oxazin-3-ylmethyl}-benzoic acid;

3-(4-Fluoro-benzyl)-6-(3-phenyl-prop-1-ynyl)-2,3-dihydro-benzo[e][1,3]oxaxin-4-one; and

6-[3-(4-Fluoro-phenyl)-prop-1-ynyl]-3-(4-methoxy-benzyl)-2,3-dihydro-benzo[e][1,3]oxaxin-4-one; or

a pharmaceutically acceptable salt thereof.

11. The compound according to Claim 8 of Formula V selected from:

4-(6-Benzylcarbamoyl-4-oxo-4H-chromen-3-ylmethyl)-benzoic acid;

4-[6-(3-Cyano-benzylcarbamoyl-4-oxo-4H-chromen-3-ylmethyl]-benzoic acid;

3-(4-Methoxy-benzyl)-4-oxo-4H-chromene-6-carboxylic acid benzyl amide; and

3-(4-Methoxy-benzyl)-4-oxo-4H-chromene-6-carboxylic acid 3-trifluoromethyl-benzyl amide; or

a pharmaceutically acceptable salt thereof.

12. The compound according to Claim 8 of Formula VII selected from:

4-[4-Oxo-6-(3-phenyl-prop-1-ynyl)-4H-chromen-3-ylmethyl]-benzoic acid;

4-{6-[3-(3,4-Dimethylphenyl-prop-1-ynyl]-4-oxo-4H-chromen-3-ylmethyl}-benzoic acid;

3-(4-Methoxy-benzyl)-6-(3-phenyl-prop-1-ynyl)-chromen-4-one; and

3-(4-Methoxy-benzyl)-6-[3-(3-methoxy-phenyl)-prop-1-ynyl]-chromen-4-one; or

a pharmaceutically acceptable salt thereof.

13. A pharmaceutical composition, comprising a compound according to Claim 1, or a pharmaceutically acceptable salt thereof, admixed with a pharmaceutically acceptable carrier, excipient, or diluent.

14. The pharmaceutical composition according to Claim 12, comprising a compound according to Claim 9, 10, 11, or 12, or a pharmaceutically acceptable

salt thereof, admixed with a pharmaceutically acceptable carrier, excipient, or diluent.

15. A method for treating arthritis, comprising administering to a patient suffering from an arthritis disease a nontoxic antiarthritic effective amount of a compound according to Claim 1, or a pharmaceutically acceptable salt thereof.

16. The method according to Claim 15, wherein the arthritis is osteoarthritis or rheumatoid arthritis.

17. The method according to Claim 16, wherein the compound administered is a compound according to Claim 9, 10, 11, or 12, or a pharmaceutically acceptable salt thereof.